

PATENT COOPERATION TREATY

PCT

From the INTERNATIONAL BUREAU

NOTIFICATION OF THE RECORDING
OF A CHANGE(PCT Rule 92bis.1 and
Administrative Instructions, Section 422)

To:

DENERLEY, Paul, Millington
AstraZeneca
Global Intellectual Property
P.O. Box 272
Mereside, Alderly Park
Macclesfield, Cheshire SK10 4GR
ROYAUME-UNI

Date of mailing (day/month/year) 19 September 2000 (19.09.00)	IMPORTANT NOTIFICATION
Applicant's or agent's file reference PHM70402/WO	
International application No. PCT/GB99/03274	International filing date (day/month/year) 04 October 1999 (04.10.99)

1. The following indications appeared on record concerning:		
<input checked="" type="checkbox"/> the applicant	<input type="checkbox"/> the inventor	<input type="checkbox"/> the agent <input type="checkbox"/> the common representative
Name and Address ASTRAZENECA UK LIMITED 15 Stanhope Gate London W1Y 6LN United Kingdom	State of Nationality GB	State of Residence GB
	Telephone No.	
	Facsimile No.	
	Teleprinter No.	
2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning:		
<input checked="" type="checkbox"/> the person	<input type="checkbox"/> the name	<input type="checkbox"/> the address <input type="checkbox"/> the nationality <input type="checkbox"/> the residence
Name and Address ASTRAZENECA AB S-151 85 Södertälje Sweden	State of Nationality SE	State of Residence SE
	Telephone No.	
	Facsimile No.	
	Teleprinter No.	
3. Further observations, if necessary:		
4. A copy of this notification has been sent to:		
<input checked="" type="checkbox"/> the receiving Office	<input type="checkbox"/> the designated Offices concerned	
<input type="checkbox"/> the International Searching Authority	<input checked="" type="checkbox"/> the elected Offices concerned	
<input checked="" type="checkbox"/> the International Preliminary Examining Authority	<input type="checkbox"/> other:	

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland	Authorized officer Marie-José Devillard
Facsimile No.: (41-22) 740.14.35	Telephone No.: (41-22) 338.83.38

BEST AVAILABLE COPY

PARENT COOPERATION TREATY

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NOTIFICATION OF THE RECORDING
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Administrative Instructions, Section 422)

To:

DENERLEY, Paul, Millington
AstraZeneca
Global Intellectual Property
P.O. Box 272
Mereside, Alderly Park
Macclesfield, Cheshire SK10 4GR
ROYAUME-UNI

Date of mailing (day/month/year) 19 September 2000 (19.09.00)	IMPORTANT NOTIFICATION
Applicant's or agent's file reference PHM70402/WO	
International application No. PCT/GB99/03274	International filing date (day/month/year) 04 October 1999 (04.10.99)

1. The following indications appeared on record concerning:	
<input type="checkbox"/> the applicant	<input type="checkbox"/> the inventor <input checked="" type="checkbox"/> the agent <input type="checkbox"/> the common representative
Name and Address DENERLEY, Paul, Millington Global Intellectual Property AstraZeneca UK Limited Mereside, Alderley Park Macclesfield Cheshire SK10 4TG United Kingdom	State of Nationality
	State of Residence
	Telephone No. (01625) 513748
	Facsimile No. (01625) 583358
2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning:	
<input type="checkbox"/> the person	<input type="checkbox"/> the name <input checked="" type="checkbox"/> the address <input type="checkbox"/> the nationality <input type="checkbox"/> the residence
Name and Address DENERLEY, Paul, Millington AstraZeneca Global Intellectual Property P.O. Box 272 Mereside, Alderly Park Macclesfield, Cheshire SK10 4GR United Kingdom	State of Nationality
	State of Residence
	Telephone No.
	Facsimile No.
3. Further observations, if necessary:	
4. A copy of this notification has been sent to:	
<input checked="" type="checkbox"/> the receiving Office	<input type="checkbox"/> the designated Offices concerned
<input type="checkbox"/> the International Searching Authority	<input checked="" type="checkbox"/> the elected Offices concerned
<input checked="" type="checkbox"/> the International Preliminary Examining Authority	<input type="checkbox"/> other:

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland Facsimile No.: (41-22) 740.14.35	Authorized officer Marie-José Devillard Telephone No.: (41-22) 338.83.38
--	---

PARENT COOPERATION TREATY

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NOTIFICATION OF THE RECORDING
OF A CHANGE(PCT Rule 92bis.1 and
Administrative Instructions, Section 422)

From the INTERNATIONAL BUREAU

To:

DENERLEY, Paul, Millington
Global Intellectual Property
AstraZeneca UK Limited
Mereside, Alderley Park
Macclesfield
Cheshire SK10 4TG
ROYAUME-UNI

Date of mailing (day/month/year)

10 May 2000 (10.05.00)

Applicant's or agent's file reference

PHM70402/WO

IMPORTANT NOTIFICATION

International application No.

PCT/GB99/03274

International filing date (day/month/year)

04 October 1999 (04.10.99)

1. The following indications appeared on record concerning:

☐

the applicant

☐

the inventor

☒

the agent

☐

the common representative

Name and Address

DENERLEY, Paul, Millington
Global Intellectual Property
AstraZeneca PLC
Mereside, Alderley Park
Macclesfield
Cheshire SK10 4TG
United Kingdom

State of Nationality

State of Residence

Telephone No.

(01625) 513748

Facsimile No.

(01625) 583358

Teleprinter No.

2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning:

☐

the person

☐

the name

☒

the address

☐

the nationality

☐

the residence

Name and Address

DENERLEY, Paul, Millington
Global Intellectual Property
AstraZeneca UK Limited
Mereside, Alderley Park
Macclesfield
Cheshire SK10 4TG
United Kingdom

State of Nationality

State of Residence

Telephone No.

(01625) 513748

Facsimile No.

(01625) 583358

Teleprinter No.

3. Further observations, if necessary:

4. A copy of this notification has been sent to:

☒

the receiving Office

☒

the designated Offices concerned

☒

the International Searching Authority

☐

the elected Offices concerned

☐

the International Preliminary Examining Authority

☐

other:

The International Bureau of WIPO
34, chemin des Colombettes
1211 Geneva 20, Switzerland

Facsimile No.: (41-22) 740.14.35

Authorized officer

Dominique DELMAS

Telephone No.: (41-22) 338.83.38

PATENT COOPERATION TREATY

PCT

NOTIFICATION OF THE RECORDING
OF A CHANGE(PCT Rule 92bis.1 and
Administrative Instructions, Section 422)

From the INTERNATIONAL BUREAU

To:

DENERLEY, Paul, Millington
Global Intellectual Property
AstraZeneca UK Limited
Mereside, Alderley Park
Macclesfield
Cheshire SK10 4TG
ROYAUME-UNIDate of mailing (day/month/year)
10 May 2000 (10.05.00)Applicant's or agent's file reference
PHM70402/WO

IMPORTANT NOTIFICATION

International application No.
PCT/GB99/03274International filing date (day/month/year)
04 October 1999 (04.10.99)

1. The following indications appeared on record concerning:

☒ the applicant ☐ the inventor ☐ the agent ☐ the common representative

Name and Address

ZENECA LIMITED
15 Stanhope Gate
London W1Y 6LN
United Kingdom

State of Nationality

GB

State of Residence

GB

Telephone No.

Facsimile No.

Teleprinter No.

2. The International Bureau hereby notifies the applicant that the following change has been recorded concerning:

☒ the person ☐ the name ☐ the address ☐ the nationality ☐ the residence

Name and Address

ASTRAZENECA UK LIMITED
15 Stanhope Gate
London W1Y 6LN
United Kingdom

State of Nationality

GB

State of Residence

GB

Telephone No.

Facsimile No.

Teleprinter No.

3. Further observations, if necessary:

4. A copy of this notification has been sent to:

☒ the receiving Office ☒ the designated Offices concerned
☒ the International Searching Authority ☐ the elected Offices concerned
☐ the International Preliminary Examining Authority ☐ other:The International Bureau of WIPO
34, chemin des Colombettes
1211 Geneva 20, Switzerland

Authorized officer

Dominique DELMAS

Facsimile No.: (41-22) 740.14.35

Telephone No.: (41-22) 338.83.38

PATENT COOPERATION TREATY

PCT

NOTIFICATION OF ELECTION

(PCT Rule 61.2)

From the INTERNATIONAL BUREAU

To:

Assistant Commissioner for Patents
United States Patent and Trademark
Office
Box PCT
Washington, D.C.20231
ETATS-UNIS D'AMERIQUE

in its capacity as elected Office

Date of mailing (day/month/year) 16 May 2000 (16.05.00)	
International application No. PCT/GB99/03274	Applicant's or agent's file reference PHM70402/WO
International filing date (day/month/year) 04 October 1999 (04.10.99)	Priority date (day/month/year) 07 October 1998 (07.10.98)
Applicant BERNSTEIN, Peter, Robert et al	

1. The designated Office is hereby notified of its election made:

☒ in the demand filed with the International Preliminary Examining Authority on:
20 April 2000 (20.04.00)

☐ in a notice effecting later election filed with the International Bureau on:

2. The election ☒ was

☐ was not

made before the expiration of 19 months from the priority date or, where Rule 32 applies, within the time limit under Rule 32.2(b).

The International Bureau of WIPO
34, chemin des Colombettes
1211 Geneva 20, Switzerland

Facsimile No.: (41-22) 740.14.35

Authorized officer

Pascal Piriou

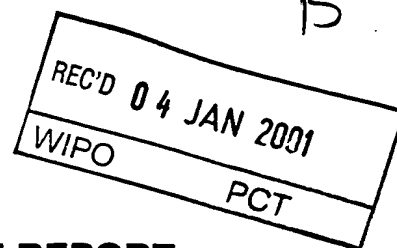
Telephone No.: (41-22) 338.83.38

PATENT COOPERATION TREATY

PCT

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

(PCT Article 36 and Rule 70)



Applicant's or agent's file reference PHM70402/WO		See Notification of Transmittal of International Preliminary Examination Report (Form PCT/IPEA/416) FOR FURTHER ACTION
International application No. PCT/GB99/03274	International filing date (day/month/year) 04/10/1999	Priority date (day/month/year) 07/10/1998
International Patent Classification (IPC) or national classification and IPC C07D211/24		
Applicant ASTRAZENECA AB et al.		

1. This international preliminary examination report has been prepared by this International Preliminary Examining Authority and is transmitted to the applicant according to Article 36.



2. This REPORT consists of a total of 6 sheets, including this cover sheet.

☐ This report is also accompanied by ANNEXES, i.e. sheets of the description, claims and/or drawings which have been amended and are the basis for this report and/or sheets containing rectifications made before this Authority (see Rule 70.16 and Section 607 of the Administrative Instructions under the PCT).

These annexes consist of a total of sheets.

3. This report contains indications relating to the following items:

- I ☒ Basis of the report
- II ☐ Priority
- III ☒ Non-establishment of opinion with regard to novelty, inventive step and industrial applicability
- IV ☐ Lack of unity of invention
- V ☒ Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement
- VI ☐ Certain documents cited
- VII ☐ Certain defects in the international application
- VIII ☐ Certain observations on the international application

Date of submission of the demand 20/04/2000	Date of completion of this report 27.12.2000
Name and mailing address of the international preliminary examining authority:  European Patent Office D-80298 Munich Tel. +49 89 2399 - 0 Tx: 523656 epmu d Fax: +49 89 2399 - 4465	Authorized officer Fanni, S Telephone No. +49 89 2399 8712 

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No. PCT/GB99/03274

I. Basis of the report

1. This report has been drawn on the basis of *(substitute sheets which have been furnished to the receiving Office in response to an invitation under Article 14 are referred to in this report as "originally filed" and are not annexed to the report since they do not contain amendments (Rules 70.16 and 70.17).)*:

Description, pages:

1-78 as originally filed

Claims, No.:

1-11 as originally filed

2. With regard to the **language**, all the elements marked above were available or furnished to this Authority in the language in which the international application was filed, unless otherwise indicated under this item.

These elements were available or furnished to this Authority in the following language: , which is:

- ☐ the language of a translation furnished for the purposes of the international search (under Rule 23.1(b)).
- ☐ the language of publication of the international application (under Rule 48.3(b)).
- ☐ the language of a translation furnished for the purposes of international preliminary examination (under Rule 55.2 and/or 55.3).

3. With regard to any **nucleotide and/or amino acid sequence** disclosed in the international application, the international preliminary examination was carried out on the basis of the sequence listing:

- ☐ contained in the international application in written form.
- ☐ filed together with the international application in computer readable form.
- ☐ furnished subsequently to this Authority in written form.
- ☐ furnished subsequently to this Authority in computer readable form.
- ☐ The statement that the subsequently furnished written sequence listing does not go beyond the disclosure in the international application as filed has been furnished.
- ☐ The statement that the information recorded in computer readable form is identical to the written sequence listing has been furnished.

4. The amendments have resulted in the cancellation of:

- ☐ the description, pages:
- ☐ the claims, Nos.:
- ☐ the drawings, sheets:

5. ☐ This report has been established as if (some of) the amendments had not been made, since they have been considered to go beyond the disclosure as filed (Rule 70.2(c)):

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No. PCT/GB99/03274

(Any replacement sheet containing such amendments must be referred to under item 1 and annexed to this report.)

6. Additional observations, if necessary:

III. Non-establishment of opinion with regard to novelty, inventive step and industrial applicability

1. The questions whether the claimed invention appears to be novel, to involve an inventive step (to be non-obvious), or to be industrially applicable have not been examined in respect of:

- ☐ the entire international application.
- ☒ claims Nos. 11.

because:

- ☒ the said international application, or the said claims Nos. 11 relate to the following subject matter which does not require an international preliminary examination (*specify*):
see separate sheet
- ☐ the description, claims or drawings (*indicate particular elements below*) or said claims Nos. are so unclear that no meaningful opinion could be formed (*specify*):
- ☐ the claims, or said claims Nos. are so inadequately supported by the description that no meaningful opinion could be formed.
- ☐ no international search report has been established for the said claims Nos. .

2. A meaningful international preliminary examination report cannot be carried out due to the failure of the nucleotide and/or amino acid sequence listing to comply with the standard provided for in Annex C of the Administrative Instructions:

- ☐ the written form has not been furnished or does not comply with the standard.
- ☐ the computer readable form has not been furnished or does not comply with the standard.

V. Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

1. Statement

Novelty (N)	Yes: Claims 1-11
	No: Claims
Inventive step (IS)	Yes: Claims
	No: Claims 1-11
Industrial applicability (IA)	Yes: Claims 1-10

**INTERNATIONAL PRELIMINARY
EXAMINATION REPORT**

International application No. PCT/GB99/03274

No: Claims

2. Citations and explanations
see separate sheet

**INTERNATIONAL PRELIMINARY
EXAMINATION REPORT - SEPARATE SHEET**

International application No. PCT/GB99/03274

ITEM III

Claim 11 relates to subject-matter considered by this Authority to be covered by the provisions of Rule 67.1(iv) PCT. Consequently, no opinion will be formulated with respect to the industrial applicability of the subject-matter of this claim (Article 34(4)(a)(i) PCT). See also the paragraph on : "Industrial applicability" in item V below.

ITEM V

Reference is made to the following documents:

D1: EP-A-0 559 538

D2: EP-A-0 630 887

D3: BIOORGANIC & MEDICINAL CHEMISTRY LETTERS, vol. 3, no. 5, 1993 ,
pages 925-930

D4: EP-A-0 515 240

D5: EP-A-0 428 434

D6: EP-A-0 474 561

NOVELTY (Article 33(2) PCT)

The present subject matter differs from D1 and D3 mainly on account of the substitution pattern on the naphthyl ring and that present residue R7 is substituted phenyl (see Table 1 from D1, lines 35-40, and compound 13, page 930 of D3).

The present subject matter differs from D2 mainly on account of the fact that the benzamide moiety in compounds of formula I according to claim 1 of D2 is replaced in the compounds according to present claim 1 by a naphthamide residue (see the definition of R4 in compounds of formula I according to claim 1 from D2 and examples 2-5, 13-22 and 25 from D2).

The present subject matter differs from D4-D6 mainly on account of the present substituents R4 and R7 (see compound 6, table 1, page 23 of D4, compound 30 from D5, and compound 46669 A, table 2, page 36 of D5).

**INTERNATIONAL PRELIMINARY
EXAMINATION REPORT - SEPARATE SHEET**

International application No. PCT/GB99/03274

INVENTIVE STEP (Article 33(3) PCT)

D2 is considered to be the closest prior art and discloses 4-(aryl-substituted)-piperidines as neurokinin receptor antagonists useful whenever such antagonism is desired, such as in the treatment of asthma and related conditions. As said above, the present subject matter differs from D2 mainly on account of the present naphthamide residue. For instance, the presence of a naphthamide instead of a benzamide residue is the only difference between example 14 from D2 and present example 1.

However, the equivalence between the phenyl and the naphthyl moiety is revealed by D3, where it is clearly states (see D3, page 928, second last paragraph), that the replacement of a phenyl with a naphthyl residue do not greatly change or reduce the in vivo activity as neurokinin receptor antagonists of the compounds described at pages 929 and 930 of D3. The present compounds are therefore considered to be an obvious solution to the problem of providing further naphthylamido-based compounds useful in the treatment of diseases where neurokinins are implicated.

Accordingly, an inventive step cannot be acknowledged for the present subject matter.

INDUSTRIAL APPLICABILITY

For the assessment of the present claim 11 on the question whether they are industrially applicable, no unified criteria exist in the PCT Contracting States. The patentability can also be dependent upon the formulation of the claims. The EPO, for example, does not recognize as industrially applicable the subject-matter of claims to the use of a compound in medical treatment, but may allow, however, claims to a known compound for first use in medical treatment and the use of such a compound for the manufacture of a medicament for a new medical treatment.

INTERNATIONAL SEARCH REPORT

International Application No

PCT/GB 99/03274

A. CLASSIFICATION OF SUBJECT MATTER

IPC 7 C07D211/24 A61K31/445 C07C255/57 C07C235/66 C07D417/12
 C07D211/26 C07F9/59 C07D413/10 C07D401/04 C07D211/22
 C07D211/34 C07D211/20

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 C07D A61K C07C C07F

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	EP 0 559 538 A (SANOFI ELF) 8 September 1993 (1993-09-08) page 21 ---	1, 10, 11
A	EP 0 630 887 A (ZENECA LTD) 28 December 1994 (1994-12-28) claim 1; examples ---	1, 10, 11
A	EMONDS-ALT X ET AL: "PHARMACOLOGICAL PROFILE AND CHEMICAL SYNTHESIS OF SR 48968, A NON -PEPTIDE ANTAGONIST OF THE NEUROKININ A (NK2) RECEPTOR" BIOORGANIC & MEDICINAL CHEMISTRY LETTERS, vol. 3, no. 5, 1 January 1993 (1993-01-01), pages 925-930, XP002068450 ISSN: 0960-894X compound 13 --- -/-	1, 10, 11

☒ Further documents are listed in the continuation of box C.

☒ Patent family members are listed in annex.

Special categories of cited documents:

- "A" document defining the general state of the art which is not considered to be of particular relevance
- "E" earlier document but published on or after the international filing date
- "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- "O" document referring to an oral disclosure, use, exhibition or other means
- "P" document published prior to the international filing date but later than the priority date claimed

- "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
- "&" document member of the same patent family

Date of the actual completion of the international search

15 December 1999

Date of mailing of the international search report

19/01/2000

Name and mailing address of the ISA

European Patent Office, P.B. 5818 Patentlaan 2
 NL - 2280 HV Rijswijk
 Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,
 Fax: (+31-70) 340-3016

Authorized officer

De Jong, B

INTERNATIONAL SEARCH REPORT

Inter. Patent Application No

PCT/GB 99/03274

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No
A	EP 0 515 240 A (SANOFI ELF) 25 November 1992 (1992-11-25) page 23; claim 1 ---	1, 10, 11
A	EP 0 428 434 A (SANOFI SA) 22 May 1991 (1991-05-22) claim 1; table I ---	1, 10, 11
A	EP 0 474 561 A (SANOFI SA) 11 March 1992 (1992-03-11) claim 1; examples 6, 29, 30 -----	1, 10, 11

INTERNATIONAL SEARCH REPORT

Information on patent family members

International Application No

PCT/GB 99/03274

Patent document cited in search report		Publication date	Patent family member(s)	Publication date
EP 0559538	A	08-09-1993	FR 2688218 A	10-09-1993
			FR 2688219 A	10-09-1993
			CA 2090785 A	04-09-1993
			WO 9318002 A	16-09-1993
			HU 70167 A	28-09-1995
			JP 6507425 T	25-08-1994
			MX 9301158 A	01-09-1993
			US 5674881 A	07-10-1997
			US 5773620 A	30-06-1998
EP 0630887	A	28-12-1994	AT 182583 T	15-08-1999
			AU 673063 B	24-10-1996
			AU 6320394 A	15-12-1994
			CA 2124048 A	25-11-1994
			CN 1098094 A	01-02-1995
			DE 69419667 D	02-09-1999
			FI 942381 A	25-11-1994
			HU 70445 A,B	30-10-1995
			IL 109734 A	24-09-1998
			IL 120895 A	24-09-1998
			JP 6340625 A	13-12-1994
			NO 941906 A	25-11-1994
			NZ 260566 A	26-07-1996
			US 5654299 A	05-08-1997
EP 0515240	A	25-11-1992	FR 2676054 A	06-11-1992
			AT 158574 T	15-10-1997
			AU 657321 B	09-03-1995
			AU 1591892 A	05-11-1992
			CA 2067924 A	04-11-1992
			CS 9201328 A	18-11-1992
			DE 69222352 D	30-10-1997
			DE 69222352 T	09-04-1998
			DK 515240 T	11-05-1998
			ES 2109987 T	01-02-1998
			FI 921950 A	04-11-1992
			GR 3025277 T	27-02-1998
			HU 65273 A,B	02-05-1994
			IL 101762 A	16-10-1996
			JP 5140103 A	08-06-1993
			MX 9202026 A	01-11-1992
			NO 178572 B	15-01-1996
			NZ 242584 A	27-04-1995
			RU 2089547 C	10-09-1997
			US 5606065 A	25-02-1997
			US 5411971 A	02-05-1995
EP 0428434	A	22-05-1991	FR 2654100 A	10-05-1991
			FR 2663329 A	20-12-1991
			AU 668018 B	18-04-1996
			AU 5924594 A	02-06-1994
			AU 649973 B	09-06-1994
			AU 6583890 A	23-05-1991
			CA 2029275 A	07-05-1991
			FI 97540 B	30-09-1996
			FI 952956 A	15-06-1995
			FI 952957 A	15-06-1995
			FI 980227 A	02-02-1998

INTERNATIONAL SEARCH REPORT

Information on patent family members

International Application No

PCT/GB 99/03274

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
EP 0428434 A		IL 96241 A	31-03-1996
		IL 111292 A	31-03-1996
		JP 3206086 A	09-09-1991
		LV 10713 A	20-06-1995
		LV 10713 B	20-10-1995
		MX 9203638 A	01-09-1992
		NO 177299 B	15-05-1995
		NO 950239 A,B,	07-05-1991
		NO 950240 A,B,	07-05-1991
		NZ 235985 A	26-10-1993
		NZ 245411 A	26-10-1993
		PL 166565 B	30-06-1995
		PL 166582 B	30-06-1995
		PT 95790 A	13-09-1991
		RU 2084453 C	20-07-1997
		US 5686609 A	11-11-1997
		US 5618938 A	08-04-1997
		US 5317020 A	31-05-1994
		PL 165758 B	28-02-1995
		PL 165854 B	28-02-1995
EP 0474561 A	11-03-1992	FR 2666335 A	06-03-1992
		FR 2678267 A	31-12-1992
		AT 174332 T	15-12-1998
		AU 657272 B	09-03-1995
		AU 8354291 A	12-03-1992
		CA 2050639 A,C	06-03-1992
		CS 9102724 A	18-03-1992
		DE 69130597 D	21-01-1999
		DE 69130597 T	20-05-1999
		ES 2127722 T	01-05-1999
		FI 914174 A,B,	06-03-1992
		GR 3029435 T	28-05-1999
		HU 9500521 A	30-10-1995
		IL 99320 A	31-07-1995
		JP 2620435 B	11-06-1997
		JP 4261155 A	17-09-1992
		LT 585 A,B	27-12-1994
		LV 10606 A	20-04-1995
		LV 10606 B	20-04-1996
		NO 177226 B	02-05-1995
		NZ 239661 A	27-06-1994
		PL 167994 B	30-12-1995
		PT 98849 A,B	31-07-1992
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PATENT COOPERATION TREATY

PCT

INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference M70402/WO	FOR FURTHER ACTION see Notification of Transmittal of International Search Report (Form PCT/ISA/220) as well as, where applicable, item 5 below.	
International application No. PCT/GB 99/ 03274	International filing date (day/month/year) 04/10/1999	(Earliest) Priority Date (day/month/year) 07/10/1998
Applicant ZENECA LIMITED et al.		

This International Search Report has been prepared by this International Searching Authority and is transmitted to the applicant according to Article 18. A copy is being transmitted to the International Bureau.

This International Search Report consists of a total of 5 sheets.

☒ It is also accompanied by a copy of each prior art document cited in this report.

1. Basis of the report

a. With regard to the **language**, the international search was carried out on the basis of the international application in the language in which it was filed, unless otherwise indicated under this item.

☐ the international search was carried out on the basis of a translation of the international application furnished to this Authority (Rule 23.1(b)).

b. With regard to any **nucleotide and/or amino acid sequence** disclosed in the international application, the international search was carried out on the basis of the sequence listing :

☐ contained in the international application in written form.

☐ filed together with the international application in computer readable form.

☐ furnished subsequently to this Authority in written form.

☐ furnished subsequently to this Authority in computer readable form.

☐ the statement that the subsequently furnished written sequence listing does not go beyond the disclosure in the international application as filed has been furnished.

☐ the statement that the information recorded in computer readable form is identical to the written sequence listing has been furnished

2. ☒ **Certain claims were found unsearchable** (See Box I).

3. ☐ **Unity of invention is lacking** (see Box II).

4. With regard to the **title**,

☐ the text is approved as submitted by the applicant.

☒ the text has been established by this Authority to read as follows:

NAPHTHALENECARBOXAMIDES AS TACHYKININ RECEPTOR ANTAGONISTS

5. With regard to the **abstract**,

☐ the text is approved as submitted by the applicant.

☒ the text has been established, according to Rule 38.2(b), by this Authority as it appears in Box III. The applicant may, within one month from the date of mailing of this international search report, submit comments to this Authority.

6. The figure of the **drawings** to be published with the abstract is Figure No. _____

☐ as suggested by the applicant.

☐ because the applicant failed to suggest a figure.

☐ because this figure better characterizes the invention.

☐ None of the figures.

INTERNATIONAL SEARCH REPORT

International application No.

GB 99/ 03274

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)

This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☒ Claims Nos.: 11
because they relate to subject matter not required to be searched by this Authority, namely:
Remark: Although claim 11
is directed to a method of treatment of the human/animal
body, the search has been carried out and based on the alleged
effects of the compounds.
2. ☐ Claims Nos.:
because they relate to parts of the International Application that do not comply with the prescribed requirements to such
an extent that no meaningful International Search can be carried out, specifically:
3. ☐ Claims Nos.:
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)

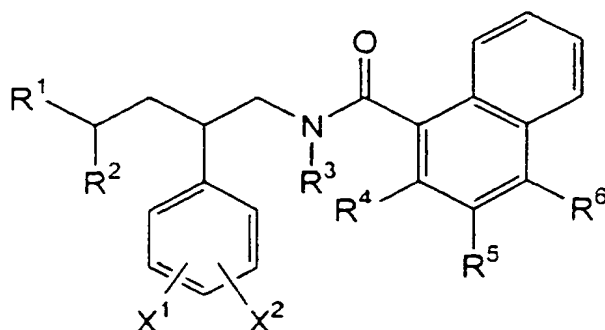
This International Searching Authority found multiple inventions in this international application, as follows:

1. ☐ As all required additional search fees were timely paid by the applicant, this International Search Report covers all
searchable claims.
2. ☐ As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment
of any additional fee.
3. ☐ As only some of the required additional search fees were timely paid by the applicant, this International Search Report
covers only those claims for which fees were paid, specifically claims Nos.:
4. ☐ No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is
restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

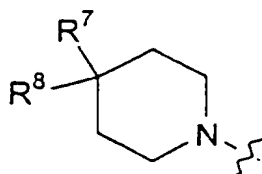
Remark on Protest

- ☐ The additional search fees were accompanied by the applicant's protest.
- ☐ No protest accompanied the payment of additional search fees.

A compound having the formula



wherein R¹ is oxo, -OR^a, -OC(=O)R^b; or



R^1 is H; or R^1 is $-OR^c$ and R^2 is $-OR^d$; or R^1 and R^2 together form $-O(CH_2)_mO-$; and any pharmaceutically-acceptable salt thereof along with their use in treating depression, anxiety, asthma, rheumatoid arthritis, Alzheimer's disease, cancer, schizophrenia, oedema, allergic rhinitis, inflammation, pain, gastrointestinal-hypermotility, anxiety, emesis, Huntington's disease, psychoses including depression, hypertension, migraine, bladder hypermotility, or urticaria, along with methods of making the compounds and pharmaceutical compositions containing the compounds.

INTERNATIONAL SEARCH REPORT

International Application No

PCT/GB 99/03274

A. CLASSIFICATION OF SUBJECT MATTER

IPC 7 C07D211/24 A61K31/445 C07C255/57 C07C235/66 C07D417/12
 C07D211/26 C07F9/59 C07D413/10 C07D401/04 C07D211/22
 C07D211/34 C07D211/20

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 C07D A61K C07C C07F

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	EP 0 559 538 A (SANOFI ELF) 8 September 1993 (1993-09-08) page 21 ---	1, 10, 11
A	EP 0 630 887 A (ZENECA LTD) 28 December 1994 (1994-12-28) claim 1; examples ---	1, 10, 11
A	EMONDS-ALT X ET AL: "PHARMACOLOGICAL PROFILE AND CHEMICAL SYNTHESIS OF SR 48968, A NON -PEPTIDE ANTAGONIST OF THE NEUROKININ A (NK2) RECEPTOR" BIOORGANIC & MEDICINAL CHEMISTRY LETTERS, vol. 3, no. 5, 1 January 1993 (1993-01-01), pages 925-930, XP002068450 ISSN: 0960-894X compound 13 --- -/--	1, 10, 11



Further documents are listed in the continuation of box C.



Patent family members are listed in annex.

* Special categories of cited documents:

"A" document defining the general state of the art which is not considered to be of particular relevance

"E" earlier document but published on or after the international filing date

"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)

"O" document referring to an oral disclosure, use, exhibition or other means

"P" document published prior to the international filing date but later than the priority date claimed

"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention

"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone

"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.

"&" document member of the same patent family

Date of the actual completion of the international search

15 December 1999

Date of mailing of the international search report

19/01/2000

Name and mailing address of the ISA

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Authorized officer

De Jong, B

INTERNATIONAL SEARCH REPORT

International Application No

PCT/GB 99/03274

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	EP 0 515 240 A (SANOFI ELF) 25 November 1992 (1992-11-25) page 23; claim 1 ---	1, 10, 11
A	EP 0 428 434 A (SANOFI SA) 22 May 1991 (1991-05-22) claim 1; table I ---	1, 10, 11
A	EP 0 474 561 A (SANOFI SA) 11 March 1992 (1992-03-11) claim 1; examples 6, 29, 30 -----	1, 10, 11

INTERNATIONAL SEARCH REPORT

Information on patent family members

International Application No

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